

§Appl. No. 09/965,807
Amdt. dated December 2, 2004
Reply to Office Action of, July 2, 2004

Listing of Claims:

Please **amend** the claims as follows:

Claim 1	(Cancelled)
Claim 2	(Cancelled)
Claim 3	(Cancelled)
Claim 4	(Cancelled)
Claim 5	(Cancelled)
Claim 6	(Cancelled)
Claim 7	(Cancelled)
Claim 8	(Cancelled)
Claim 9	(Cancelled)
Claim 10	(Cancelled)
Claim 11	(Cancelled)
Claim 12	(Cancelled)
Claim 13	(Cancelled)
Claim 14	(Cancelled)
Claim 15	(Cancelled)
Claim 16	(Cancelled)
Claim 17	(Cancelled)
Claim 18	(Cancelled)
Claim 19	(Cancelled)
Claim 20	(Cancelled)
Claim 21	(Cancelled)

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Claim 22 (Currently Amended) An isolated naturally-occurring mutant human aspartoacylase polypeptide ~~which~~ having either an altered ability to hydrolyze N-acetyl-aspartic acid to aspartate and acetate, as compared with a ~~normal~~ wild-type human aspartoacylase, or incapable of hydrolyzing N-acetyl-aspartic acid to aspartate and acetate, and ~~having~~ comprising the amino acid sequence SEQ ID NO: 2 of wild-type human aspartoacylase, except for said mutation, which is

E285 > A,

Y231 > X, and/or

A305 > E,

~~or an allelic variant of said mutant aspartoacylase~~ a naturally-occurring mutant allele of said wild-type human aspartoacylase.

Claim 23 (Cancelled)

Claim 24 (Previously Presented) A mutant aspartoacylase of claim 22, wherein the glutamic acid at amino acid position 285 is substituted by alanine.

Claim 25 (Cancelled)

Claim 26 (Cancelled)

Claim 27 (Cancelled)

Claim 28 (Cancelled)

Claim 29 (Cancelled)

Claim 30 (Cancelled)

Claim 31 (Cancelled)

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Claim 32	(Cancelled)
Claim 33	(Cancelled)
Claim 34	(Cancelled)
Claim 35	(Cancelled)
Claim 36	(Cancelled)
Claim 37	(Cancelled)
Claim 38	(Cancelled)
Claim 39	(Cancelled)
Claim 40	(Cancelled)
Claim 41	(Cancelled)
Claim 42	(Cancelled)
Claim 43	(Cancelled)
Claim 44	(Cancelled)
Claim 45	(Cancelled)
Claim 46	(Cancelled)
Claim 47	(Cancelled)
Claim 48	(Cancelled)
Claim 49	(Cancelled)
Claim 50	(Cancelled)
Claim 51	(Cancelled)
Claim 52	(Cancelled)
Claim 53	(Cancelled)
Claim 54	(Cancelled)
Claim 55	(Cancelled)
Claim 56	(Cancelled)
Claim 57	(Cancelled)

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Claim 58 (Cancelled)

Claim 59 (Cancelled)

Claim 60 (Cancelled)

Claim 61 (Cancelled)

Claim 62 (Cancelled)

Claim 63 (Previously Presented) A transgenic mouse exhibiting Canavan disease symptoms.

Claim 64 (Cancelled)

Claim 65 (Cancelled)

Claim 66 (Currently Amended) A fragment of a mutant human aspartoacylase of claim 22, comprising an aspartoacylase epitope which is immunologically-effective to elicit antibodies that selectively bind to said human aspartoacylase.

Claim 67 (Currently Amended) A recombinant ~~normal~~ wild-type human aspartoacylase capable of hydrolyzing N-acetyl aspartic acid to aspartate and acetate, ~~having~~ comprising an amino acid sequence which has a sequence identity of at least 95% to the sequence of SEQ ID NO: 2.

Claim 68 (Currently Amended) A fragment of a recombinant ~~normal~~ wild-type human

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aspartoacylase of claim ~~20~~ 67, comprising an aspartoacylase epitope which is immunologically-effective to elicit antibodies that selectively bind to said human aspartoacylase.

Claim 69 (Currently Amended) A pharmaceutical composition, comprising an isolated ~~normal~~ wild-type human aspartoacylase ~~having~~ comprising the amino acid sequence SEQ ID NO: 2, or a naturally-occurring polymorphic form thereof, and a pharmaceutically acceptable carrier.

Claim 70 (Currently Amended) An isolated normal human aspartoacylase ~~having~~ comprising the amino acid sequence SEQ ID NO: 2, or a naturally-occurring polymorphic form thereof, which is free of other cellular components.

Claim 71 (Currently Amended) An isolated ~~normal~~ normal-type human aspartoacylase ~~having~~ comprising the amino acid sequence SEQ ID NO: 2, or a naturally-occurring polymorphic form thereof, which is free of other human proteins.

Claim 72 (Currently Amended) A preparation which consists essentially of a ~~normal~~ wild-type human aspartoacylase ~~having~~ comprising the amino acid sequence SEQ ID NO: 2, or a naturally-occurring polymorphic form thereof.

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Claim 73 (Currently Amended) An isolated ~~normal~~ wild-type human aspartoacylase ~~having~~ comprising the amino acid sequence SEQ ID NO: 2, or a naturally-comprising polymorphic form thereof, in a concentration which can be administered to a patient at a dosage of 0.1 to 100 U/kg.

Claim 74 (Currently Amended) A ~~normal~~ wild-type human aspartoacylase ~~having~~ comprising the amino acid sequence SEQ ID NO: 2, or a naturally-occurring polymorphic form thereof, produced by

(a) culturing a host cell transformed with a vector comprising a DNA which encodes for a normal human aspartoacylase of claim ~~20~~ 80 in a cell culture medium under conditions whereby the aspartoacylase is expressed, and

(b) isolating the thus-produced ~~normal~~ wild-type aspartoacylase.

Claim 75 (Currently Amended) A ~~normal~~ wild-type human aspartoacylase ~~having~~ comprising the amino acid sequence SEQ ID NO: 2, or a naturally-occurring polymorphic form thereof, produced in a bacterium, a fungus, or a non-human mammalian cell.

Claim 76 (Withdrawn) An immunologically active anti-aspartoacylase polyclonal or monoclonal antibody specific for an aspartoacylase polypeptide of claim 20.

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Claim 77 **(Withdrawn)** An immunologically active anti-aspartoacylase polyclonal or monoclonal antibody specific for an aspartoacylase polypeptide of claim 22.

Claim 78 **(Withdrawn)** A hybridoma producing a monoclonal antibody specific for an aspartoacylase polypeptide of claim 20.

Claim 79 **(Withdrawn)** A hybridoma producing a monoclonal antibody specific for an aspartoacylase polypeptide of claim 22.

Claim 80 **(Currently Amended)** A recombinant ~~normal~~ wild-type human aspartoacylase capable of hydrolyzing N-acetyl aspartic acid to aspartate and acetate, having comprising the amino acid sequence SEQ ID NO: 2, or a naturally-occurring polymorphic form thereof.

Claim 81 **(Currently Amended)** A ~~normal~~ wild-type human aspartoacylase polypeptide purified to homogeneity and capable of hydrolyzing N-acetyl-aspartic acid to aspartate and acetate.

Claim 82 **(Previously Presented)** The aspartoacylase of claim 81 having SEQ ID NO: 2.

Claim 83 **(New)** An isolated polypeptide of claim 22 which is encoded by a nucleic acid

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which specifically hybridizes under stringent conditions to a nucleotide sequence of SEQ ID
NO:1.

Claim 84 (New) An isolated polypeptide of claim 66 which is encoded by a nucleic acid
which specifically hybridizes under stringent conditions to a nucleotide sequence of SEQ ID
NO:1.

Claim 85 (New) An isolated polypeptide of claim 67 which is encoded by a nucleic acid
which specifically hybridizes under stringent conditions to a nucleotide sequence of SEQ ID
NO:1.

Claim 86 (New) An isolated polypeptide of claim 68 which is encoded by a nucleic acid
which specifically hybridizes under stringent conditions to a nucleotide sequence of SEQ ID
NO:1.

Claim 87 (New) An isolated polypeptide of claim 71 which is encoded by a nucleic acid
which specifically hybridizes under stringent conditions to a nucleotide sequence of SEQ ID
NO:1.

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Claim 88 **(New)** An isolated polypeptide of claim 72 which is encoded by a nucleic acid which specifically hybridizes under stringent conditions to a nucleotide sequence of SEQ ID NO:1.